## Yale University New Haven, Connecticut 06508

SCHOOL OF MEDICINE

ABRAHAM RIBICOFF RESEARCH FACILITIES

Department of Psychiatry and Connecticut Mental Health Center 34 Park Street

February 9, 1989

203-789-7151

Dr. Marshall Nirenberg
National Institute of Heart, Lung
and Blood
Building 36, RC27
9000 Rockville Pike
Bethesda, MD 20892

Dear Dr. Nirenberg:

Based on your earlier work on neuroblastoma cells, we have been studying the involvement of the cyclic AMP system in opiate addiction in the rat locus coeruleus. Similar to your earlier findings, we have found that chronic morphine treatment increases levels of certain G-proteins, adenylate cyclase activity, cyclic AMP-dependent protein kinase activity, and certain phosphoprotein substrates for the protein kinase in this brain region. Interestingly, these effects were all specific to the locus coeruleus, as they were not observed in the other brain regions examined. Enclosed please find reprints that describe some of our work on opiate action in the locus coeruleus. I thought you might be interested in them, as they support and extend your novel proposal of a role for the cyclic AMP system in mediating opiate addiction elaborated over ten years ago.

Sincerely.

Eric J. Nestler, M.D., Ph.D.

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Assistant Professor of Psychiatry

and Pharmacology

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